

**ABSTRACT**

**[000215]** The present invention provides, *inter alia*, methods of selecting a single-stranded oligomeric compounds for inhibiting RNA expression, methods of generating double-stranded oligomeric compounds, methods of identifying optimized double-stranded oligomeric compounds, methods of selecting optimized single-stranded oligomeric compounds, methods of selecting optimized double-stranded oligomeric compounds, methods of identifying multifunctional oligomeric compounds, methods for optimizing target region selection for modulation of RNA expression, methods of optimizing expression modulation of RNA, and the like. The present invention further provides oligomeric compounds, 8-80 nucleobases in length targeted to a target RNA, wherein said oligomeric compound hybridizes to said target RNA and inhibits RNA levels by at least 50% in both single-stranded and double-stranded forms, and multifunctional oligomeric compounds.